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ATTORNEY'S DOCKET NUMBER TRANSMITTAL LETTER TO THE UNITED STATES 3190-088 DESIGNATED/ELECTED OFFICE (DO/EO/US) see 37 CFR 1.5) CONCERNING A FILING UNDER 35 U.S.C. 371 INTERNATIONAL APPLICATION NO. INTERNATIONAL FILING DATE PRIORITY DATE CLAIMED September 30, 2004 September 30, 2003 PCT/JP2004/014378 TITLE OF INVENTION: INHIBITION OF NERVE CELL DEATH BY INHIBITING DEGRADATION OF SHC3, ATF6 OR CREBL1 BY HTRA2 AND METHOD OF AMELIORATING NEURODEGENERATIVE DISEASES APPLICANT(S) FOR DO/EO/US: Hirofumi DOI and Ken SAITO Applicant herewith submits to the United States Designated/Elected Office (DO/EO/US) the following items and other information: 1. | x | This is a FIRST submission of items concerning a filing under 35 U.S.C. 371. This is a **SECOND** or **SUBSEQUENT** submission of items concerning a filing under 35 U.S.C. 371. This is an express request to begin national examination procedures (35 U.S.C. 371 (f)). The submission must include items (5), (6), (9) and (21) indicated below. 4. x The US has been elected (Article 31). 5. X A copy of the International Application as filed (35 U.S.C. 371 (c)(2)) a. x is attached hereto (required only if not communicated by the International Bureau). b. | x | has been communicated by the International Bureau. is not required, as the application was filed in the United States Receiving Office (RO/US). 6. | x | An English language translation of the International Application as filed (35 U.S.C. 371 (c)(2)). a. | x | is attached hereto. has been previously submitted under 35 U.S.C. 154(d)(4). 7. X Amendments to the claims of the International Application under PCT Article 34 (35 U.S.C. 371(c)(3)) are attached hereto (required only if not communicated by the International Bureau). have been communicated by the International Bureau. have not been made; however, the time limit for making such amendments has NOT expired. have not been made and will not be made. An English language translation of the amendments to the claims under PCT Article 19 (35 U.S.C. 371(c)(3)) An oath or declaration of the inventor(s) (35 U.S.C. 371(c)(4)). An English language translation of the annexes to the International Preliminary Examination Report under PCT 10. Article 36 (35 U.S.C. 371(c)(5)). Items 11 to 20 below concern document(s) or information included: An Information Disclosure Statement under 37 CFR 1.97 and 1.98. An assignment document for recording. A separate cover sheet in compliance with 37 CFR 3.28 and 3.31 is included. 13. x | A preliminary amendment. An Application Data Sheet A substitute specification. 15. A power of attorney and/or address change letter. A computer-readable form of the sequence listing in accordance with PCT Rule 13ter.2 and 35 U.S.C. 1.821 – 1.825. A second copy of the published international application under 35 U.S.C. 154(d)(4). A second copy of the English language translation of the international application under 35 U.S.C. 154(d)(4). Other items or information: PCT/ISA/237 (in Japanese)

This collection of information is required by 37 CFR 1.53(b). The information is required to obtain or retain a benefit by the public which is to file (and by the USPTO to process) an application. Confidentiality is governed by 35 U.S.C. 122 and 37 CFR 1.14. This collection is estimated to take 12 minutes to complete, including gathering, preparing, and submitting the completed application form to the USPTO. Time will vary depending upon the individual case. Any comments on the amount of time you require to complete this form and/or suggestions for reducing this burden, should be sent to the Chief Information Officer, U.S. Patent and Trademark Office, U.S. Department of Commerce, P.O. Box 1450, Alexandria, VA 22313-1450. DO NOT SEND FEES OR COMPLETED FORMS TO THIS ADDRESS. SEND TO: Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450. If you need assistance in completing the form, call 1-800-PTO-9199 and select option 2

Request for the recording a change submitted December 1, 2005 (in Japanese)

PCT/IB/308 PCT/IB/304

## IAP6 Rec'd PCT/PTO 29 DEC 2005

PTO-1390 (Rev. 12-2004)
Approved for use through 03/31/2007 OMB 0651-0021
U.S. Patent and Trademark Office; U.S. DEPARTMENT OF COMMERCE

Under the Paperwork Reduction Act of 1995, no persons are required to respond to a collection of information unless it displays a valid OMB control number. 5629 INTERNATIONAL APPLICATION NO. PCT/JP2004/014378 U.S. APPLICATION NO. (if kg ATTORNEY'S DOCKET NUMBER 3190-088 Unknown 21. X The following fees are submitted: 冈 a) Basic national fee \$300.00 \$300.00  $\boxtimes$ b) Examination fee .......\$200.00 \$ 200.00 冈 c) Search fee ......\$500.00 \$ 500.00 TOTAL OF ABOVE CALCULATIONS = \$ 1,000.00 \$1,000.00 Additional fee for specification and drawings filed in paper over 100 sheets (excluding sequence listing or computer program listing filed in an electronic medium). The fee is \$250.00 for each additional 50 sheets of paper or fraction thereof. Number of each additional 50 or fraction **Total Sheets** Extra sheets thereof (round up to a whole number) /50 =x \$250.00 \$ 0.00 49 - 100 =Surcharge of \$130.00 for furnishing the oath or declaration later than Months from the earliest claimed \$130.00 priority date (37 CFR 1.492(e)). **CLAIMS** NUMBER FILED NUMBER EXTRA RATE 12 - 20 =0 \$50.00 \$ 0.00 Total claims х \$ 200.00 4 - 3 =\$200.00 Independent claims 1 X MULTIPLE DEPENDENT CLAIM(S) (if applicable) \$360.00 \$ 0.00 TOTAL OF ABOVE CALCULATIONS \$1,330.00 Applicant claims small entity status. See 37 CFR 1.27. The fees indicated above are \$ 0.00 reduced by 1/2. SUBTOTAL \$ 1,330.00 Processing fee of \$130.00 for furnishing the English translation later than months from the earliest claimed priority date (37 CFR 1.492(f)). \$ 0.00 TOTAL NATIONAL FEE = \$1,330.00 Fee for recording the enclosed assignment (37 CFR 1.21(h)). The assignment must be \$ 0.00 accompanied by an appropriate cover sheet (37 CFR 3.28, 3.31). \$40.00 per property TOTAL FEES ENCLOSED \$1,130.00 Amount to be: \$ Refunded \$ Amount to be Charged to cover the above fees is enclosed. A check in the amount of \$ in the amount of \$ \_\_\_\_\_ to cover the above fees. Please charge my Deposit Account No. b. A duplicate copy of this sheet is enclosed. The Commissioner is hereby authorized to charge any additional fees which may be required, or credit any overpayment c. to Deposit Account No. 50-0925. Fees are to be charged to a credit card. WARNING: Information on this form may become public. Credit card d. information should not be included on this form. Provide credit card information and authorization on PTO-2038. NOTE: Where an appropriate time limit under 37 CFR 1.494 or 1.495 has not been met, a petition to revive (37 CFR 1.137(a) or (b)) must be filed and granted to restore the application to pending status. SEND ALL CORRESPONDENCE TO KILYK & BOWERSOX, P.L.L.C. 400 Holiday Court, Suite 102 Warrenton, VA 20186 Phone (540) 428-1701 - Facsimile (540) 428-1720 REGISTRATION NUMBER Form PTO-1390 (REV 12-2001) Date: December 29, 2005 Label No. EV790042093US I hereby certify that, on the date indicated above, I deposited this paper with identified attachments and/or fee with the U.S. Postal Service and that it was addressed for delivery to the Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450 by "Express Mail Post Office to Addressee" service. NUM Kim Blum

Name (Print) Signature